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PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

CERTIFICATE OF MAILING

I hereby certify that this **INFORMATION DISCLOSURE STATEMENT** and documents submitted therewith are being deposited with the United States Postal Service as first class mail, postage prepaid thereon, in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on the date indicated below.

Ariel Fletcher
Ariel Fletcher

6/24/2004
Date

Applicant: Madison)
) Group: 1652
Serial No.: 10/705,633)
) Confirmation No: 4805
Filed: November 10, 2003)
) Examiner: Unassigned
For: TISSUE TYPE PLASMINOGEN)
ACTIVATOR (t-PA) VARIANTS:) Our Ref.: TSRI 568.1 Div 1
COMPOSITIONS AND METHODS)
OF USE)

INFORMATION DISCLOSURE STATEMENT

Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

Dear Sir:

In recognition of its continuing duty to disclose pursuant to 37 CFR §1.56, Applicant hereby submits the present Information Disclosure Statement in compliance therewith.

Pursuant to 37 CFR §1.98(d), enclosed herewith is a copy of 1) the Information Disclosure Statement and 2) the accompanying PTO Form 1449 filed in application serial no. 09/600,985. Application 09/600,985 was filed on November 13, 2000, and has issued as U.S. Patent No. 6,706,504, to which the present application is related by division. Also, pursuant to 37 CFR §1.98(d), copies of the above references are not submitted

herewith, as they are available in the related patent application 09/600,985.

Applicants understand that the interpretation given to each reference may differ from one individual to another. The PTO is therefore encouraged to independently examine the disclosed references. While the references provided in this Information Disclosure Statement may be material pursuant to 37 CFR §1.56, it shall not be construed to be an admission that the cited information is, or is considered to be, material to patentability unless specifically designated as such.

Applicants are filing the present statement pursuant to 37 CFR §1.97(b) insofar as this statement is being filed within three months of the filing of the application and/or before the mailing date of a first Office Action.

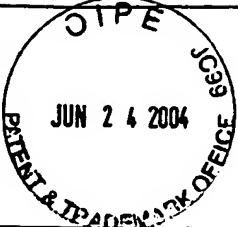
Also, in accordance with 37 CFR §1.97(g), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made or, that if made, any search was complete or exhaustive, or that no other material information as defined in 37 CFR §1.56 exists.

Respectfully submitted,

June 21, 2004
Date

Thomas Fitting
Thomas Fitting, Reg. No. 34,163

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FORM PTO-1449 U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE INFORMATION DISCLOSURE STATEMENT BY APPLICANT		ATTY DOCKET NO. TSRI 568.1	SERIAL NO. 09/600,985
		APPLICANT Madison	
		FILING DATE November 13, 2000	GROUP

U.S. PATENT DOCUMENTS

EXAM. INITIALS	DOCUMENT NUMBER	DATE	NAME	CLASS	SUB- CLASS	FILING DATE
	5,616,486	4/1/97	Anderson, et al.			

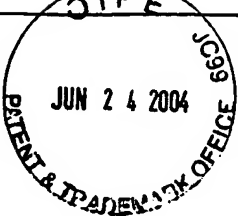
FOREIGN PATENT DOCUMENTS

EXAM. INITIALS	DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB- CLASS	TRANSLATION YES NO
	EP 0 351 246	1/17/90	EPO			
	WO 90/02798	3/22/90	PCT			
	EP 0 199 574	10/29/86	EPO			
	WO 88/05081	7/14/88	PCT			
	EP 0 241 209	10/14/87	EPO			
	EP 0 201 153	11/12/86	EPO			

OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages)

	1	Petersen, et al., Quenching of the amidolytic activity of one-chain tissue-type plasminogen activator by mutation of lysine-416, 1990, <i>Biochemistry</i> , 29(14):3451-3457.
	2	Tachias, et al., Variants of tissue-type plasminogen activator which display substantially enhanced stimulation by fibrin, 1995, <i>J. Biol. Chem.</i> , 270(31):18319-18322.
	3	Krause, et al., Properties of molecular variants of tissue-type plasminogen activator, 1989, <i>Arzneim.-Forsch.</i> , 39:632-637.
	4	van Zonneveld, et al., Mapping of epitopes on human tissue-type plasminogen activator with recombinant deletion mutant proteins, 1987, <i>Thrombosis & Haemostasis</i> , 57(1):82-86.
	5	Browne, et al., A tissue-type plasminogen activator mutant with prolonged clearance in vivo. Effect of removal of the growth factor domain, 1988, <i>J. Biol. Chem.</i> , 263:1599-1602.
	6	Madison, et al., Serpin resistant mutants of human tissue-type plasminogen activator, 1989, <i>Nature</i> , 339:721-724.
	7	Higgins, et al., The effect of the one-chain to two-chain conversion in tissue plasminogen activator: Characterization of mutations at position 275, 1990, <i>Thrombosis Res.</i> , 57:527-539.
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EXAMINER		DATE CONSIDERED

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to Applicant.

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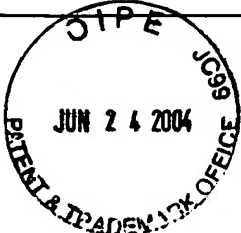
FOREIGN PATENT DOCUMENTS

EXAM. INITIALS	DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB- CLASS	TRANSLATION YES NO
	EP 0 233 013	8/19/87	EPO			
	EP 0 292 009	11/23/88	EPO			
	EP 0 293 936	12/7/88	EPO			
	EP 0 293 934	12/7/88	EPO			
	WO 88/10119	12/29/88	PCT			
	EP 0 225 286	6/10/87	EPO			

OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages)

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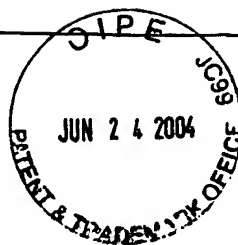
EXAM. INITIALS		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB- CLASS	TRANSLATION YES NO
		WO 87/04722	8/13/87	PCT			
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		EP 0 299 706	1/18/89	EPO			
		EP 0 238 304	9/23/87	EPO			
		EP 0 231 624	8/12/87	EPO			

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FORM PTO-1449 U.S. DEPARTMENT OF COMMERCE
PATENT AND TRADEMARK OFFICE



ATTY DOCKET NO.
TSRI 568.1

SERIAL NO.
09/600,985

APPLICANT
Madison

FILING DATE
November 13, 2000

GROUP

INFORMATION DISCLOSURE
STATEMENT BY APPLICANT

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EXAM. INITIALS	DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB- CLASS	TRANSLATION YES NO
	EP 0 290 118	11/9/88	EPO			
	EP 0 240 334	10/7/87	EPO			

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PATENT AND TRADEMARK OFFICE



ATTY DOCKET NO.
TSR1 568.1 D1

SERIAL NO.
10705,633

APPLICANT
Madison

FILING DATE
11/20/2003

GROUP
1652

INFORMATION DISCLOSURE
STATEMENT BY APPLICANT

U.S. PATENT DOCUMENTS

EXAM. INITIALS		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB- CLASS	FILING DATE
	1	5,520,913	05/1996	Anderson et al.			
	2	5,525,477	06/1996	Hassouna			
	3	5,246,850	09/1993	Bennett et al.			

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	4	Lamba, et al., "The 2.3 A Crystal Structure of the Catalytic Domain of Recombinant Two-chain Human Tissue-type Plasminogen Activator", <u>J. Mol. Biol.</u> 258: 117-135 (1996)
	5	Strandberg, et al., "Variants of Tissue-type Plasminogen Activator with Substantially Enhanced Response and Selectivity toward Fibrin Co-factors", <u>J. Biol. Chem.</u> 270(40): 23444-23449 (1995)
	6	Tate, et al., "Functional Role of Proteolytic Cleavage at Arginine-275 of Human Tissue Plasminogen Activator as Assessed by Site-Directed Mutagenesis", <u>Biochemistry</u> 26: 338-343 (1987)
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